EXHIBIT D

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1. Affitoxin--a novel recombinant, HER2-specific, anticancer agent for targeted therapy of HER2-positive tumors.

Zielinski R, Lyakhov I, Jacobs A, Chertov O, Kramer-Marek G, Francella N, Stephen A, Fisher R, Blumenthal R, Capala J.

J Immunother. 2009 Oct;32(8):817-25. PMID: 19752752 [PubMed - in process]

2. An internalizing antibody specific for the human asialoglycoprotein receptor.

Trahtenherts A. Benhar I.

Hybridoma (Larchmt). 2009 Aug;28(4):225-33. PMID: 19663694 [PubMed - indexed for MEDLINE]

3. Tumor-targeted PE38KDEL delivery via PEGylated anti-HER2 immunoliposomes.

Gao J, Zhong W, He J, Li H, Zhang H, Zhou G, Li B, Lu Y, Zou H, Kou G, Zhang D, Wang H, Guo Y, Zhong Y.

Int J Pharm, 2009 Jun 5;374(1-2):145-52. Epub 2009 Mar 24.

PMID: 19446771 [PubMed - indexed for MEDLINE]

Recombinant immunotoxins containing truncated bacterial toxins for the treatment of hematologic malignancies.

Kreitman RJ.

BioDrugs. 2009;23(1):1-13. doi: 10.2165/00063030-200923010-00001. Review.

PMID: 19344187 [PubMed - indexed for MEDLINE] Free PMC Article Free text

 Combination therapy using three novel prolactin receptor antagonist-based fusion proteins effectively inhibits tumor recurrence and metastasis in HER2/neu transgenic mice.

Tomblyn S, Springs AE, Langenheim JF, Chen WY.

Int J Oncol. 2009 Apr;34(4):1139-46.

PMID: 19287973 [PubMed - indexed for MEDLINE]

 Treatment of hepatocellular carcinoma in mice with PE38KDEL type I mutant-loaded poly(lactic-co-glycolic acid) nanoparticles conjugated with humanized SM5-1 F(ab') fragments.

Gao J, Kou G, Chen H, Wang H, Li B, Lu Y, Zhang D, Wang S, Hou S, Qian W, Dai J, Zhao J, Zhong Y, Guo Y.

Mol Cancer Ther. 2008 Oct;7(10):3399-407.

PMID: 18852143 [PubMed - indexed for MEDLINE] Free Article

7. Antitumor activity of immunotoxins with T-cell receptor-like specificity against human melanoma xenografts.

Klechevsky E, Gallegos M, Denkberg G, Palucka K, Banchereau J, Cohen C, Reiter Y.

Cancer Res. 2008 Aug 1;68(15):6360-7.

PMID: 18676861 [PubMed - indexed for MEDLINE] Free PMC Article Free text

8. PE38KDEL-loaded anti-HER2 nanoparticles inhibit breast tumor progression with reduced toxicity and immunogenicity.

Gao J, Kou G, Wang H, Chen H, Li B, Lu Y, Zhang D, Wang S, Hou S, Qian W, Dai J, Zhao J, Zhong Y, Guo Y.

Breast Cancer Res Treat. 2009 May;115(1):29-41. Epub 2008 May 15.

PMID: 18481173 [PubMed - indexed for MEDLINE]

Preparation and characterization of PE38KDEL-loaded anti-HER2 nanoparticles for targeted cancer therapy.

Chen H, Gao J, Lu Y, Kou G, Zhang H, Fan L, Sun Z, Guo Y, Zhong Y.

J Control Release, 2008 Jun 24;128(3):209-16. Epub 2008 Mar 19.

PMID: 18450313 [PubMed - indexed for MEDLINE]

Expression, purification, and characterization of an immunotoxin containing a humanized anti-CD25 single-chain fragment variable antibody fused to a modified truncated Pseudomonas exotoxin A.

Wang H, Dai J, Li B, Fan K, Peng L, Zhang D, Cao Z, Qian W, Wang H, Zhao J, Guo Y.

Protein Expr Purif. 2008 Mar;58(1):140-7. Epub 2007 Sep 19.

PMID: 17950620 [PubMed - indexed for MEDLINE]

11. Administration of a CD25-directed immunotoxin, LMB-2, to patients with metastatic melanoma induces a selective partial reduction in regulatory T cells in vivo.

Powell DJ Jr, Felipe-Silva A, Merino MJ, Ahmadzadeh M, Allen T, Levy C, White DE, Mavroukakis S, Kreitman RJ, Rosenberg SA, Pastan I. J Immunol. 2007 Oct 1;179(7):4919-28.

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12. EGFRvIII-targeted immunotoxin induces antitumor immunity that is inhibited in the absence of CD4+ and CD8+ T cells.

Ochiai H, Archer GE, Herndon JE 2nd, Kuan CT, Mitchell DA, Bigner DD, Pastan IH, Sampson JH.

Cancer Immunol Immunother. 2008 Jan; 57(1):115-21. Epub 2007 Jul 19.

PMID: 17634939 [PubMed - indexed for MEDLINE] Free PMC Article Free text

13. Treatment of hepatocellular carcinoma in a mouse xenograft model with an immunotoxin which is engineered to eliminate vascular leak

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Cancer immunol immunother. 2007 Nov;56(11):1775-83. Epub 2007 Apr 13.

PMID: 17431617 [PubMed - indexed for MEDLINE]

14. Immunotoxins in the treatment of refractory hairy cell leukemia.

Kreitman RJ, Pastan I.

Hematol Oncol Clin North Am. 2006 Oct;20(5):1137-51, viii. Review.

PMID: 16990113 [PubMed - indexed for MEDLINE]

15. Selective elimination of human regulatory T lymphocytes in vitro with the recombinant immunotoxin LMB-2.

Attia P, Powell DJ Jr, Maker AV, Kreitman RJ, Pastan I, Rosenberg SA.

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PMID: 16531821 [PubMed - indexed for MEDLINE] Free PMC Article Free text

16. Rapid downregulation of beta-actin-based CAG promoter and filamentous actin in injured podocytes.

Asano T, Matsusaka T, Mizutani S, Ichikawa I.

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17. CLL immunotoxins.

Frankel AE, Kreitman RJ.

Leuk Res. 2005 Sep;29(9):985-6. Epub 2005 Mar 17. PMID: 16038723 [PubMed - indexed for MEDLINE]

18. Preparation and characterization of fusion protein truncated Pseudomonas Exotoxin A (PE38KDEL) in Escherichia coli.

Song S, Xue J, Fan K, Kou G, Zhou Q, Wang H, Guo Y.

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PMID: 15922623 [PubMed - indexed for MEDLINE]

19. Genetic engineering of glomerular sclerosis in the mouse via control of onset and severity of podocyte-specific injury.

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J Am Soc Nephrol. 2005 Apr;16(4):1013-23. Epub 2005 Mar 9.

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Biochemistry (Mosc). 2005 Jan;70(1):62-8.

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21. Hodgkin lymphoma therapy with interleukin-4 receptor-directed cytotoxin in an infiltrating animal model.

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22. Interleukin-4 receptor cytotoxin as therapy for human malignant pleural mesothelioma xenografts.

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Ann Thorac Surg. 2004 Aug;78(2):436-43; discussion 436-43.

PMID: 15276492 [PubMed - indexed for MEDLINE]

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Methods Find Exp Clin Pharmacol. 2004 Apr;26(3):211-44.

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24. Interleukin-4-Pseudomonas exotoxin chimeric fusion protein for malignant glioma therapy.

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25. Multidrug-resistant tumor cells remain sensitive to a recombinant interleukin-4-Pseudomonas exotoxin, except when overexpressing the multidrug resistance protein MRP1.

de Jong MC, Scheffer GL, Broxterman HJ, Hooijberg JH, Slootstra JW, Meloen RH, Kreitman RJ, Husain SR, Joshi BH, Puri RK, Scheper RJ

Clin Cancer Res. 2003 Oct 15;9(13):5009-17.

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Wang JS, Wang SM, Chen GR, Yang DJ.

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Clin Cancer Res. 2002 Nov;8(11):3503-11.

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Br J Cancer. 2002 Jan 21;86(2):285-91.

PMID: 11870521 [PubMed - indexed for MEDLINE] Free PMC Article Free text

32. Sensitization of B-cell chronic lymphocytic leukemia cells to recombinant immunotoxin by immunostimulatory phosphorothioate oligodeoxynucleotides.

Decker T, Hipp S, Kreitman RJ, Pastan I, Peschel C, Licht T.

Blood. 2002 Feb 15;99(4):1320-6.

PMID: 11830482 [PubMed - indexed for MEDLINE] Free Article

33. Interleukin 4 receptor-directed cytotoxin therapy for human head and neck squamous cell carcinoma in animal models.

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Clin Cancer Res. 2002 Jan;8(1):281-6.

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34. Internalization property of interleukin-4 receptor alpha chain increases cytotoxic effect of interleukin-4 receptor-targeted cytotoxin in cancer cells.

Kawakami K, Kawakami M, Leland P, Puri RK.

Clin Cancer Res. 2002 Jan;8(1):258-66.

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35. Biologic therapy for psoriasis: a brief history, I.

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Cutis. 2001 Nov;68(5):331-6. Review.

PMID: 11766118 [PubMed - indexed for MEDLINE]

36. In situ expression of interleukin-4 (IL-4) receptors in human brain tumors and cytotoxicity of a recombinant IL-4 cytotoxin in primary glioblastoma cell cultures.

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Methods Mol Biol. 2001;166:155-76. Review. No abstract available.

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39. Quantification of immunotoxin number for complete therapeutic response.

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PMID: 11217362 [PubMed - indexed for MEDLINE]

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Onda M, Willingham M, Wang QC, Kreitman RJ, Tsutsumi Y, Nagata S, Pastan I.

J Immunol. 2000 Dec 15;165(12):7150-6.

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41. Human breast carcinoma cells express type II IL-4 receptors and are sensitive to antitumor activity of a chimeric IL-4-Pseudomonas exotoxin fusion protein in vitro and in vivo.

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43. Structure, function, and targeting of interleukin 4 receptors on human head and neck cancer cells.

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44. Cytotoxic activity of disulfide-stabilized recombinant immunotoxin RFB4(dsFv)-PE38 (BL22) toward fresh malignant cells from patients with B-cell leukemias.

Kreitman RJ, Margulies I, Stetler-Stevenson M, Wang QC, FitzGerald DJ, Pastan I.

Clin Cancer Res. 2000 Apr;6(4):1476-87.

PMID: 10778980 [PubMed - indexed for MEDLINE] Free Article

45. Phase I trial of recombinant immunotoxin anti-Tac(Fv)-PE38 (LMB-2) in patients with hematologic malignancies.

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J Biol Chem. 2000 Mar 17;275(11):7566-73.

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Robbins DH, Margulies I, Stetler-Stevenson M, Kreitman RJ.

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PMID: 10690555 [PubMed - indexed for MEDLINE] Free Article

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Kuska B

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Husain SR, Kreitman RJ, Pastan I, Puri RK.

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PMID: 10395315 [PubMed - indexed for MEDLINE]

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Skrepnik N, Zieske AW, Bravo JC, Gillespie AT, Hunt JD.

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56. Recombinant heregulin-Pseudomonas exotoxin fusion proteins: interactions with the heregulin receptors and antitumor activity in vivo. Yang D, Kuan CT, Payne J, Kihara A, Murray A, Wang LM, Alimandi M, Pierce JH, Pastan I, Lippman ME.

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57. Recombinant toxins containing human granulocyte-macrophage colony-stimulating factor and either pseudomonas exotoxin or diphtheria toxin kill gastrointestinal cancer and leukemia cells.

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Interleukin-4 receptor expression on AIDS-associated Kaposi's sarcoma cells and their targeting by a chimeric protein comprised of Interleukin-4 receptor expression on AIDG-associates in a circularly permuted interleukin-4 and Pseudomonas exotoxin.

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59. Preclinical development of a recombinant toxin containing circularly permuted interleukin 4 and truncated Pseudomonas exotoxin for therapy of malignant astrocytoma.

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Cancer Res. 1996 Dec 15;56(24):5631-7.

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Mavroudis DA, Jiang YZ, Hensel N, Lewalle P, Couriel D, Kreitman RJ, Pastan I, Barrett AJ.

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Kuan CT, Pastan I.

Biochemistry. 1996 Mar 5;35(9):2872-7.

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66. Activity of anti-erbB-2 recombinant toxin OLX-209 on lung cancer cell lines in the absence of erbB-2 gene amplification.

Kasprzyk PG, Sullivan TL, Hunt JD, Gubish CT, Scoppa CA, Oelkuct M, Bird R, Fischer PH, Siegfried JM, King CR.

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68. Cytotoxic activity of chimeric toxins containing the epidermal growth factor-like domain of heregulins fused to PE38KDEL, a truncated recombinant form of Pseudomonas exotoxin.

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76. Recombinant immunotoxins containing the VH or VL domain of monoclonal antibody B3 fused to Pseudomonas exotoxin.

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Search: pe40kdel

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Xi Y, Yuan Z, Zhang H, Guan H, Kong F, Liu N, Liang F, Cui J, Guo S, Sun Y, Xi C.

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